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Sequence Listing

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Alfred L. George, Jr.

<120> PURIFIED AND ISOLATED POTASSIUM-CHLORIDE COTRANSPORTER NUCLEIC ACIDS AND POLYPEPTIDES AND THERAPEUTIC AND SCREENING METHODS USING SAME

<130> Attorney Docket No. 1242-26-2

<140> US/09/835,976A

<141> 2001-04-16

<160> 131

<170> PatentIn Ver. 2.1

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gag ggc ccc gag ccc gag cgc ccc agc ccg gga gat gga aat cca aga 145 Glu Gly Pro Glu Pro Glu Arg Pro Ser Pro Gly Asp Gly Asn Pro Arg 35 40 45

gaa aac agc cca ttc ntc aac aat gtc gag gtg gaa caa gag agc ttc 193
Glu Asn Ser Pro Phe Xaa Asn Asn Val Glu Val Glu Glu Glu Ser Phe
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						atc Ile		673
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aac atg cca ggt cct ccc aaa aac cgg cag gga gac gag aac tac atg Asn Met Pro Gly Pro Pro Lys Asn Arg Gln Gly Asp Glu Asn Tyr Met 1040 1045 1050 1055
gag ttt ctt gaa gtc ctg acc gag ggg ctg aac aga gtc ctc ctg gtc 3217 Glu Phe Leu Glu Val Leu Thr Glu Gly Leu Asn Arg Val Leu Leu Val 1060 1065 1070
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Asn Ser Pro Phe Xaa Asn Asn Val Glu Val Glu Glu Ser Phe Phe
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Glu Gly Lys Asn Met Ala Leu Phe Glu Glu Glu Met Asp Ser Asn Pro 65 70 75 80

Met Val Ser Ser Leu Xaa Asn Lys Leu Ala Asn Tyr Thr Asn Leu Ser 85 90 95

Gln Gly Val Val Glu His Glu Glu Asp Glu Glu Ser Arg Arg Glu 105 100 Ala Lys Ala Pro Arg Met Gly Thr Phe Ile Gly Val Tyr Leu Pro Cys 120 Leu Gln Asn Ile Leu Gly Val Ile Leu Phe Leu Arg Leu Thr Trp Ile 135 Val Gly Val Ala Gly Val Leu Glu Ser Phe Leu Ile Val Ala Met Cys 150 155 Cys Thr Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala Ile Ala Thr Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Tyr Met Ile Ser Arg Ser 185 Leu Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys Phe Tyr Leu Gly Thr Thr Phe Ala Gly Ala Met Tyr Ile Leu Gly Thr Ile Glu Ile Phe 215 Leu Thr Tyr Ile Ser Pro Gly Ala Ala Ile Phe Gln Ala Glu Ala Ala 230 Gly Glu Ala Ala Ala Met Leu His Asn Met Arg Val Tyr Gly Thr 245 250 Cys Thr Leu Val Leu Met Ala Leu Val Val Phe Val Gly Val Lys Tyr Val Asn Lys Leu Ala Leu Val Phe Leu Ala Cys Val Val Leu Ser Ile 280 Leu Ala Ile Tyr Ala Gly Val Ile Lys Ser Ala Phe Asp Pro Pro Asp Ile Pro Val Cys Leu Leu Gly Asn Arg Thr Leu Ser Arg Arg Ser Phe Asp Ala Cys Val Lys Ala Tyr Gly Ile His Asn Asn Ser Ala Thr Ser Ala Leu Trp Gly Leu Phe Cys Asn Gly Ser Gln Pro Ser Ala Ala Cys Asp Glu Tyr Phe Ile Gln Asn Asn Val Thr Glu Ile Gln Gly Ile Pro 355 360 Gly Ala Ala Ser Gly Val Phe Leu Glu Asn Leu Trp Ser Thr Tyr Ala His Ala Gly Ala Phe Val Glu Lys Lys Gly Val Pro Ser Val Pro Val 395 390 Ala Glu Glu Ser Arg Ala Ser Thr Leu Pro Tyr Val Leu Thr Asp Ile 405 410 415

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Asp Lys His Met Glu Ala Gln Arg Ala Glu Glu Asn Ile Arg Ser Leu 745 Met Ser Thr Glu Lys Thr Lys Gly Phe Cys Gln Leu Val Val Ser Ser 760 Ser Leu Arg Asp Gly Met Ser His Leu Ile Gln Ser Ala Gly Leu Gly 775 Gly Leu Lys His Asn Thr Val Leu Met Ala Trp Pro Ala Ser Trp Lys 795 Gln Glu Asp Asn Pro Phe Ser Trp Lys Asn Phe Val Asp Thr Val Arg Asp Thr Thr Ala Ala His Gln Ala Leu Leu Val Ala Lys Asn Val Asp Ser Phe Pro Gln Asn Gln Glu Arg Phe Gly Gly His Ile Asp Val Trp Trp Ile Val His Asp Gly Gly Met Leu Met Leu Pro Phe Leu 855 Leu Arg Gln His Lys Val Trp Arg Lys Cys Arg Met Arg Ile Phe Thr Val Ala Gln Val Asp Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Gln 885 890 Met Phe Leu Tyr His Leu Arg Ile Ser Ala Glu Val Glu Val Val Glu 900 Met Val Glu Asn Asp Ile Ser Ala Phe Thr Tyr Glu Arg Thr Leu Met 920 Met Glu Gln Arg Ser Gln Met Leu Lys Gln Met Gln Leu Ser Lys Asn Glu Gln Glu Arg Glu Ala Gln Leu Ile His Asp Arg Asn Thr Ala Ser His Thr Ala Ala Ala Ala Arg Thr Gln Ala Pro Pro Thr Pro Asp Lys Val Gln Met Thr Trp Thr Arg Glu Lys Leu Ile Ala Glu Lys Tyr Arg Ser Arg Asp Thr Ser Leu Ser Gly Phe Lys Asp Leu Phe Ser Met Lys Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala Val Lys Leu Asn 1015 Gly Val Val Xaa Asn Lys Ser Gln Asp Ala Gln Leu Val Leu Leu Asn 1025 1030 1035 Met Pro Gly Pro Pro Lys Asn Arg Gln Gly Asp Glu Asn Tyr Met Glu 1045 1050 1055

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					gca Ala											656
					atg Met 170											704
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cta ccc a Leu Pro												1520
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tta tct Leu Ser '				_	_							1856

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						aac Asn										1952
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	cac His												2960
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	agt Ser												3104
	agg Arg												3152
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Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Met Ser 50 55 60

Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Pro Ser Asp 65 70 75 80

Arg Thr Ser His Pro Gln Asp Val Ile Glu Asp Asp Gly His Lys Lys 85 90 95

Ala Arg Asn Ala Tyr Leu Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu
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Tyr Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Met Asp Thr Arg
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Pro Lys Val Ser Ser Leu Leu Asn Arg Met Ala Asn Tyr Thr Asn Leu 130 135 140

Lys Lys Lys Pro Thr Lys Thr Pro Gln Met Gly Thr Phe Met Gly Val 165 170 175

Tyr Leu Pro Cys Leu Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Arg 180 185 190

Leu Thr Trp Val Val Gly Thr Ala Gly Val Leu Gln Ala Phe Ala Ile 195 200 205

Val Leu Ile Cys Cys Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser 210 215 220

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Ile Glu Ile Phe Leu Val Tyr Ile Val Pro Arg Ala Ala Ile Phe His 275 280 285 .-

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Val	Thr	Gly	Ile	Met 485	Ala	Gly	Ser	Asn	Arg 490	Ser	Gly	Asp	Leu	Lys 495	Asp
Ala	Gln	Lys	Ser 500	Ile	Pro	Ile	Gly	Thr 505	Ile	Leu	Ala	Ile	Leu 510	Thr	Thr
Ser	Phe	Val 515	Tyr	Leu	Ser	Asn	Val 520	Val	Leu	Phe	Gly	Ala 525	Cys	Ile	Glu
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Pro	Arg	Leu	Leu 580	Gln	Ala	Ile	Ala	Lys 585	Asp	Asn	Ile	Ile	Pro 590	Phe	Leu
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Phe Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp 930 935 940

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Val Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr 965 970 975

Leu Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser 980 985 990

Lys Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser 995 1000 1005

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Thr Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met 1025 1030 1035 1040

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Leu Asn Met Arg Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala 1060 1065 1070

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Val Leu Leu Asn Met Pro Gly Pro Pro Arg Asn Pro Glu Gly Asp Glu 1090 1095 1100

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cgg (912
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atc (Ile (320																1008
gta (1056
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gcc A	_			_	_											1392

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ctg (1920
ctc 1 Leu 1 640				_	_	_	_	_		_	_	-		1968
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Lys Ala Gly Lys Gly Leu Thr Ile Val Gly Ser Val Ile Val Gly Asn 775 Phe Leu Glu Asn Tyr Gly Asp Ala Leu Ala Ala Glu Gln Thr Ile Lys 790 795 His Leu Met Glu Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val 805 810 Ala Ala Lys Leu Lys Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly 825 Leu Gly Gly Met Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly 840 Trp Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr Val Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn 870 Val Ser Phe Pro Ser Asn Val Glu Gln Phe Ser Glu Gly Asn Ile 890 885 Asp Val Arg Trp Ile Val His Asp Gly Gly Met Leu Met Leu Pro 900 Phe Leu Leu Lys Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile 920 Phe Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp 935 Leu Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val 950 955 Val Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr 970 Leu Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser Lys Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser Met Leu Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu 1015 Thr Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met 1025 1030 1035 Ala Ser Arg Gly Gln Lys Val Lys Ser Met Glu Gly Phe Gln Asp Leu 1045 1050 Leu Asn Met Arg Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala 1060 1065

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1085

1080

1075

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Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Leu Ser 50 55 60

Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Ser Ser Asp
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Arg Thr Ser Asn Pro Gln Asp Val Thr Glu Asp Pro Ser Gln Asn Ser 85 90 95

Ile Thr Gly Glu His Ser Gln Leu Leu Asp Asp Gly His Lys Lys Ala
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Arg Asn Ala Tyr Leu Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu Tyr 115 120 125

Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Glu Met Asp Thr Arg Pro 130 135 140

Lys Val Ser Ser Leu Leu Asn Arg Met Ala Asn Tyr Thr Asn Leu Thr 145 150 155 160

Gln Gly Ala Lys Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly Lys 165 170 175

Lys Lys Pro Thr Lys Ser Pro Gln Met Gly Thr Phe Met Gly Val Tyr 180 185 190

Leu Pro Cys Leu Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Arg Leu 195 200 205

Thr Trp Val Val Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Ile Val 210 220

Leu Ile Cys Cys Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala 225 230 235 240

Ile Ala Thr Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Phe Met Ile 245 250 255

Ser Arg Ala Leu Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys Phe 265 Tyr Leu Gly Thr Thr Phe Ala Ala Ala Met Tyr Ile Leu Gly Ala Ile 280 Glu Ile Phe Leu Val Tyr Ile Val Pro Arg Ala Ala Ile Phe Arg Ser 295 300 Asp Asp Ala Leu Lys Glu Ser Ala Ala Met Leu Asn Asn Met Arg Val 310 315 Tyr Gly Thr Ala Phe Leu Val Leu Met Val Leu Val Val Phe Ile Gly 325 330 Val Arg Tyr Val Asn Lys Phe Ala Ser Leu Phe Leu Ala Cys Val Ile 340 Val Ser Ile Leu Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe Ala 360 Pro Pro His Phe Pro Val Cys Met Leu Gly Asn Arg Thr Leu Ser Ser 370 Arg His Leu Asp Ile Cys Ser Lys Thr Lys Glu Val Asp Asn Met Thr 395 390 Val Pro Ser Lys Leu Trp Gly Phe Phe Cys Asn Ser Ser Gln Phe Phe Asn Ala Thr Cys Asp Glu Tyr Phe Val His Asn Asn Val Ile Ser Ile 425 Gln Gly Ile Pro Gly Leu Ala Ser Gly Ile Ile Thr Glu Asn Leu Trp Ser Asn Tyr Leu Pro Lys Gly Glu Ile Ile Glu Lys Pro Ser Ala Lys 455 Ser Ser Asp Val Leu Gly Asn Leu Asn His Glu Tyr Val Leu Ala Asp 470 Ile Thr Thr Ser Phe Thr Leu Leu Val Gly Ile Phe Phe Pro Ser Val 490 Thr Gly Ile Met Ala Gly Ser Asn Arg Ser Gly Asp Leu Lys Asp Ala Gln Lys Ser Ile Pro Ile Gly Thr Ile Leu Ala Ile Leu Thr Thr Ser 520 Phe Val Tyr Leu Ser Asn Val Val Leu Phe Gly Ala Cys Ile Glu Gly 530 535 Val Val Leu Arg Asp Lys Phe Gly Asp Ala Val Lys Gly Asn Leu Val 550 Val Gly Thr Leu Ser Trp Pro Ser Pro Trp Val Ile Val Ile Gly Ser 570

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Val Arg Trp Ile Val His Asp Gly Gly Met Leu Met Leu Pro Phe 915 920 925

Leu Leu Lys Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile Phe 930 935 940

Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp Leu 945 950 955 960

Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val 965 970 975

Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr Leu 980 985 990

Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser Lys 995 1000 1005

Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser Met 1010 1015 1020

Leu Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu Thr 1025 1030 1035 1040

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Ser Arg Gly Gln Lys Val Lys Ser Met Glu Gly Phe Gln Asp Leu Leu 1060 1065 1070

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ccg agt cag aac tcc atc aca ggg gag cac agc cag ctg tta gat gac 256 Pro Ser Gln Asn Ser Ile Thr Gly Glu His Ser Gln Leu Leu Asp Asp 45 50 55
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							cac His 320									1072
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ttc :			_	-									1456
gat (Pro					1504
atc (1552
gca i													1600
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atc q													1696
ntc a Xaa 1													1744
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acc f		_				_	_		_		_		1840
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atg 1	_					_	_	_	_	_			1936
cga a Arg '													1984
tct (Ser)													2032

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Glu His Ser Gln Leu Leu Asp Asp Gly His Lys Lys Ala Arg Asn Ala 50 55 60

Tyr Xaa Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu Tyr Phe Asp Lys
65 70 75 80

Asn Leu Ala Leu Phe Glu Glu Glu Met Asp Thr Arg Pro Lys Val Ser

Leu	Leu	Asn 100	Arg	Met	Ala	Asn	Tyr 105	Thr	Asn	Leu	Thr	Gln 110	Gly	Ala
Glu	His 115	Glu	Glu	Ala	Glu	Asn 120	Ile	Thr	Glu	Gly	Lys 125	Lys	Lys	Pro
Lys 130	Ser	Pro	Gln	Met	Gly 135	Thr	Phe	Met	Gly	Val 140	Tyr	Leu	Pro	Cys
Gln	Asn	Ile	Phe	Gly 150	Val	Ile	Leu	Phe	Leu 155	Arg	Leu	Thr	Trp	Val 160
Gly	Thr	Ala	Gly 165	Ile	Leu	Gln	Ala	Phe 170	Ala	Ile	Val	Leu	Ile 175	Cys
Cys	Cys	Thr 180	Met	Leu	Thr	Ala	Ile 185	Ser	Met	Ser	Ala	Ile 190	Ala	Thr
Gly	Val 195	Val	Pro	Ala	Gly	Gly 200	Ser	Tyr	Phe	Met	Ile 205	Ser	Arg	Ala
Gly 210	Pro	Glu	Phe	Gly	Gly 215	Ala	Val	Gly	Leu	Cys 220	Phe	Tyr	Leu	Gly
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Leu Asp	Trp 355 Glu	340 Gly Tyr	Lys Phe Phe	Phe Val	Cys His 375	Glu Asn 360 Asn	Val 345 Ser Asn	330 Asp Ser Val	Asn Gln Ile	Met Phe Ser 380	Thr Phe 365	350 Asn Gln	335 Pro Ala Gly	Ser Thr
	Lys 130 Gln Gly Cys Gly 210 Thr Val Lys Phe Asn 290 Ala	Lys Ser 130 Ser 130 Ser 130 Ser 130 Ser 130 Ser 140 Se	Glu His Glu Lys Ser Pro 130 Gln Asn Ile Gly Thr Ala Cys Cys Thr 180 Gly Val Val 195 Gly Pro Glu 210 Thr Phe Ala Val Tyr Ile Lys Glu Ser 260 Phe Leu Val 275 Asn Lys Phe 290 Ala Ile Tyr	Glu His Glu Glu Lys Ser Pro Gln 130 Gln Asn Ile Phe Gly Thr Ala Gly 165 Cys Cys Thr Met 180 Gly Pro Glu Pro 210 Thr Phe Ala Ala Val Tyr Ile Val 245 Lys Glu Ser Ala 260 Phe Leu Val Leu 275 Ala Ile Tyr Ala	GluHis 115GluGluAlaLysSerProGlnMetGlnAsnIlePheGly 150GlyThrAlaGlyIleCysCysThrMetLeuGlyValValProAlaGlyProGluPheGlyThrPheAlaAlaAlaValTyrIleValProLysGluSerAlaAlaPheLeuValLeuMetAsnLysPheAlaSerAlaIleTyrAlaGlyAlaIleTyrAlaGly	GluHis 115GluGluAlaGluLysSerProGlnMetGly 135GlnAsnIlePheGly 150ValGlyThrAlaGlyIleLeuCysCysThrMetLeuThrGlyValProAlaGlyGlyProGluProAlaGlyThrPheAlaAlaAlaMetValTyrIleValProArgLysGluSerAlaAlaMetPheLeuValLeuMetValAsnLysPheAlaSerLeuAlaIleTyrAlaGlyAlaAlaIleTyrAlaGlyAla	GluHis 115GluGluAlaGluAsn 120Lys GlnSer ProGlnMet 135Thr 135Thr 136GlnAsnIlePhe 150ValIleGlyThr 180Met ProLeuThr AlaGlyVal 195Pro GluPhe 210Gly 215AlaThrPhe 210AlaAla 245MetTyrValTyrIle 260Val 245Pro ArgAlaLysGlu 260Ser 260AlaAlaMetLeuPhe 275Val 280Leu 280MetLeu 280AlaIleTyrAlaGly 310AlaIle	Glu His Glu Glu Ala Glu Asn Ile Lys Ser Pro Gln Met Gly Thr Phe Gln Asn Ile Phe Gly Val Ile Leu Gly Thr Ala Gly Ile Leu Gln Ala Cys Cys Thr Met Leu Thr Ala Ile Gly Val Pro Ala Gly Gly Ser Gly Pro Glu Phe Gly Gly Ala Val Thr Phe Ala Ala Ala Ala Met Tyr Ile Val Tyr Ile Val Pro Arg Ala Ala Lys Glu Ser Ala Ala Met Leu Asn Lys Phe Leu Met Val Leu Asn 280 Phe Leu Met Val Leu 280 Val 2	Glu His Glu Glu Ala Glu Asn Ile Thr Lys Ser Pro Gln Met Gly Thr Phe Met Gln Asn Ile Phe Gly Val Ile Leu Phe Gly Thr Ala Gly Ile Leu Gln Ala Phe Cys Thr Ala Gly Ile Leu Gln Ala Phe Gly Val Val Pro Ala Gly Gly Ser Tyr Gly Pro Glu Phe Gly Gly Ala Val Gly Thr Phe Ala Ala Ala Met Tyr Ile Leu Val Tyr Ile Val Pro Arg Ala Ala Ala Phe Leu Val Leu Ala Met Leu Ala Ala Lys Phe Ala Ala Ser Leu P	Glu His Glu Glu Ala Glu Asn Ile Thr Glu Lys Ser Pro Gln Met Gly Thr Phe Met Gly Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Gly Yal Val Pro Ala Gly Ser Tyr Phe Gly Yal Val Phe Gly Gly Ser Tyr Phe Gly Pro Glu Phe Gly Gly Ser Tyr Phe Gly Pro Ala Ala Ala Ala Val Val Leu Gly Leu L	Glu His 115 Glu Glu Ala Glu Asn 120 Ile Thr Glu Gly Val 140 Lys Ser Pro Gln Met Gly Thr Phe Met Gly Thr Phe Leu Ala Ile Leu Arg 155 Arg Arg 140 Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Ile Gly Thr Ala Gly Ile Leu Thr Ala Ile Ser Met Ser Gly Yal Val Pro Ala Gly Ala Val Phe Met Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys Thr Phe Ala Ala Ala Ala Ala Ala Ile Phe Arg Lys Glu Ser Ala Ala Met Tyr Ile Leu Phe	Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly Lys 125 Lys Ser Pro Gln Met Gly Yal Ile Leu Phe Leu Arg Leu 150 Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Arg Leu 165 Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Ile Ser Met Ser Ala 180 Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala 195 Gly Val Val Pro Ala Gly Gly 200 Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys Phe 200 Thr Phe Ala Ala Ala Ala Met Tyr Ile Leu Gly Ala Ile 235 Val Tyr Ile Val Pro Arg Ala Ala Ile Phe Arg Ser Lys Glu Ser Ser Net Arg Val 265 Lys Glu Ser Ala Ala Met Leu Asn Asn Met Arg Val 265 Asn Lys Phe Ala Ser Leu 295 Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe Ala 300 Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe Ala	Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly Lys Lys <td>Glu His Glu Ala Glu Ass Ile Thr Glu Gly Lys Lys</td>	Glu His Glu Ala Glu Ass Ile Thr Glu Gly Lys Lys

Val Leu Gly Asn Leu Asn His Glu Tyr Val Leu Ala Asp Ile Thr Thr 425 Ser Phe Thr Leu Leu Val Gly Ile Phe Phe Pro Ser Val Thr Gly Ile 440 Met Ala Gly Ser Asn Arg Ser Gly Asp Leu Lys Asp Ala Gln Lys Ser 455 Ile Pro Ile Gly Thr Ile Leu Ala Ile Leu Thr Thr Ser Phe Val Tyr 470 475 Leu Ser Asn Val Val Leu Phe Gly Ala Cys Ile Glu Gly Val Val Leu 485 490 Arg Asp Lys Phe Gly Asp Ala Val Lys Gly Asn Leu Val Val Gly Thr Leu Ser Trp Pro Ser Pro Trp Val Ile Val Ile Gly Ser Phe Phe Ser 520 Thr Cys Gly Ala Gly Leu Gln Ser Xaa Thr Gly Ala Pro Arg Leu Leu 535 Gln Ala Ile Ala Lys Asp Asn Ile Ile Pro Phe Leu Arg Val Phe Gly 555 550 His Ser Lys Ala Asn Gly Glu Pro Thr Trp Ala Leu Leu Leu Thr Ala . 570 Ala Ile Ala Glu Leu Gly Ile Leu Ile Ala Ser Leu Asp Leu Val Ala Pro Ile Leu Ser Met Phe Phe Leu Met Cys Tyr Leu Phe Val Asn Leu 600 Ala Cys Ala Leu Gln Thr Leu Leu Arg Thr Pro Asn Trp Arg Pro Arg 610 Phe Arg Tyr Tyr His Trp Ala Leu Ser Phe Met Gly Met Ser Ile Cys Leu Ala Leu Met Phe Ile Ser Ser Trp Tyr Tyr Ala Ile Val Ala Met Val Ile Ala Gly Met Ile Tyr Lys Tyr Ile Glu Tyr Gln Gly Ala Glu Lys Glu Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Ser Ala Ala Arg 680 Phe Ala Leu Leu Arg Leu Glu Glu Gly Pro Pro His Thr Lys Asn Trp Arg Pro Gln Leu Leu Val Leu Lys Leu Asp Glu Asp Leu His Val 710 715 Lys His Pro Arg Leu Leu Thr Phe Ala Ser Gln Leu Lys Ala Gly Lys 725 730 735

Gly Leu Thr Ile Val Gly Ser Val Ile Val Gly Asn Phe Leu Glu Asn 745 740 Tyr Gly Asp Ala Leu Ala Ala Glu Gln Thr Ile Lys His Leu Met Glu 760 Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val Ala Ala Lys Leu 775 Lys Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly Leu Gly Met 795 Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly Trp Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr Val Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn Val Ser Phe Phe 840 835 Pro Ser Asn Val Glu Gln Phe Ser Glu Gly Asn Ile Asp Val Arg Trp 855 Ile Val His Asp Gly Gly Met Leu Met Leu Pro Phe Leu Leu Lys 870 Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile Phe Thr Val Ala 890 885 Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Ala Thr Phe 905 900 Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val Glu Met His 920 Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr Leu Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser Lys Thr Glu Arg 955 Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser Met Xaa Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu Thr Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met Ala Ser Arg Gly 1000 Gln Lys Val Lys Ser Met Glu Gly Phe Gln Asp Leu Leu Asn Met Arg 1015 Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala Val Lys Leu Asn 1025 1030 1035 Glu Val Ile Val Asn Lys Ser His Glu Ala Lys Leu Val Leu Leu Asn

1050

1055

1045

Met Pro Gly Pro Pro Arg Asn Pro Glu Gly Asp Glu Asn Tyr Met Glu 1060 1065 1070

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Gly Gly Ser Glu Val Ile Thr Ile Tyr Ser 1090 1095

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gca gaa aac aat gag ggt gga aaa aag aag ccg gtg cag gcc cca cgc

Ala Glu Asn Asn Glu Gly Gly Lys Lys Pro Val Gln Ala Pro Arg

90

85

288

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	_				_									gca Ala		384
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	_	_			_	_	_		_	_			_	gtg Val		480
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gcc Ala	atg Met	tac Tyr 195	atc Ile	ctg Leu	ggc Gly	acc Thr	atc Ile 200	gaa Glu	atc Ile	ctg Leu	ctg Leu	gct Ala 205	tac Tyr	ctc Leu	ttc Phe	624
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														ggc Gly		960
														ttc Phe 335		1008

cga aad Arg Asr	a Asn V														1056
ctc ato Leu Ile															1104
gtg gag Val Gli 370	a Arg S														1152
atc gad Ile Asp 385			lis E												1200
acc cto		Val G													1248
ggt tct Gly Ser	Asn A														1296
act ggo Thr Gly															1344
tcc gtt Ser Val 450	. Val I														1392
aag ttt Lys Phe 465			la V												1440
tgg cca Trp Pro		Pro I													1488
ggg gct Gly Ala	Gly I	ctg c Leu G 500	ag a Sln S	agc Ser	ctc Leu	acg Thr	202 203 333	gcc Ala	cca Pro	cgc Arg	ctg Leu	ctg Leu 510	cag Gln	gcc Ala	1536
atc tcg Ile Ser															1584
aag gco Lys Ala 530	Asn (1632
tgc gag Cys Glu 545			le I												1680
ctc tct Leu Sei	_	Phe P		_	_	_		_				_	_	_	1728

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	cgg aag tgc Arg Lys Cys					
	agc atc cag Ser Ile Gln 870	Met Lys Ly				
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	gct tac acc Ala Tyr Thr 900		s Thr Leu	Val Met		
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	agt atc aca Ser Ile Thr					
_	aac acg cgg Asn Thr Arg 950	_		_		_
	gaa gag aag Glu Glu Lys 965					_
	ccc agc tgc Pro Ser Cys 980		r Ser Pro	Ser Pro		
	gaa ggg gag Glu Gly Glu				Leu Thr	
	aag tcg gtg Lys Ser Val					
	ggc atc aag Gly Ile Lys 103	Asp Phe Ph		Lys Pro		
	cag tcc aac Gln Ser Asn 1045					Leu

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aac atg cct ggg cct ccc cgc aac cgc aat ggt gat gaa aac tac atg 3264 Asn Met Pro Gly Pro Pro Arg Asn Arg Asn Gly Asp Glu Asn Tyr Met 1075 1080 1085
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Phe Cys Ser Ser Arg Phe Leu Asn Ala Thr Cys Asp Glu Tyr Phe Thr Arg Asn Asn Val Thr Glu Ile Gln Gly Ile Pro Gly Ala Ala Ser Gly 345 Leu Ile Lys Glu Asn Xaa Trp Ser Ser Tyr Leu Thr Lys Gly Val Ile 360 Val Glu Arg Ser Gly Met Thr Ser Val Gly Leu Ala Asp Gly Thr Pro 375 Ile Asp Met Asp His Pro Tyr Val Phe Ser Asp Met Thr Ser Tyr Phe 390 Thr Leu Leu Val Gly Ile Tyr Phe Pro Ser Val Thr Gly Ile Met Ala 410 Gly Ser Asn Arg Ser Gly Asp Leu Arg Asp Ala Gln Lys Ser Ile Pro Thr Gly Thr Ile Leu Ala Ile Ala Thr Thr Ser Ala Val Tyr Ile Ser 440 Ser Val Val Leu Phe Gly Ala Cys Ile Glu Gly Val Val Leu Arg Asp 450 Lys Phe Gly Glu Ala Val Asn Gly Asn Leu Val Val Gly Thr Leu Ala 475 470 Trp Pro Ser Pro Trp Val Ile Val Ile Gly Ser Phe Phe Ser Thr Cys 485 490 Gly Ala Gly Leu Gln Ser Leu Thr Gly Ala Pro Arg Leu Leu Gln Ala 505 Ile Ser Arg Asp Gly Ile Val Pro Phe Leu Gln Val Phe Gly His Gly 520 515 Lys Ala Asn Gly Glu Pro Thr Trp Ala Leu Leu Thr Ala Cys Ile 535 Cys Glu Ile Gly Ile Leu Ile Ala Ser Leu Asp Glu Val Ala Pro Ile 545 550 555 Leu Ser Met Phe Phe Leu Met Cys Tyr Met Phe Val Asn Leu Ala Cys 570 Ala Val Gln Thr Leu Leu Arg Thr Pro Asn Trp Arg Pro Arg Phe Arg Tyr Tyr His Trp Thr Leu Ser Phe Leu Gly Met Ser Leu Cys Leu Ala Leu Met Phe Ile Cys Ser Trp Tyr Tyr Ala Leu Val Ala Met Leu Ile 615 Ala Gly Leu Ile Tyr Lys Tyr Ile Glu Tyr Arg Gly Ala Glu Lys Glu 630 635 625

Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Ser Ala Ala Arg Tyr Ala 650 Leu Leu Arg Leu Glu Glu Gly Pro Pro His Thr Lys Asn Trp Arg Pro 665 Gln Leu Leu Val Leu Val Arg Val Asp Gln Asp Gln Asn Val Val His 680 Pro Gln Leu Leu Ser Leu Thr Ser Gln Leu Lys Ala Gly Lys Gly Leu 695 Thr Ile Val Gly Ser Val Leu Glu Gly Thr Phe Leu Glu Asn His Pro 710 Gln Ala Gln Arg Ala Glu Glu Ser Ile Arg Arg Leu Met Glu Ala Glu 730 Lys Val Lys Gly Phe Cys Gln Val Val Ile Ser Ser Asn Leu Arg Asp Gly Val Ser His Leu Ile Gln Ser Gly Gly Leu Gly Gly Leu Gln His Asn Thr Val Leu Val Gly Trp Pro Arg Asn Trp Arg Gln Lys Glu Asp His Gln Thr Trp Arg Asn Phe Ile Glu Leu Val Arg Glu Thr Thr Ala 790 795 Gly His Leu Ala Leu Leu Val Thr Lys Asn Val Ser Met Phe Pro Gly Asn Pro Glu Arg Phe Ser Glu Gly Ser Ile Asp Val Trp Trp Ile Val 825 His Asp Gly Gly Met Leu Met Leu Leu Pro Phe Leu Leu Arg His His 835 Lys Val Trp Arg Lys Cys Lys Met Arg Ile Phe Thr Val Ala Gln Met 855 Asp Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Thr Thr Phe Leu Tyr 875 His Leu Arg Ile Thr Ala Glu Val Glu Val Glu Met His Glu Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Lys Thr Leu Val Met Glu Gln Arg Ser Gln Ile Xaa Lys Gln Met His Leu Thr Lys Asn Glu Arg Glu Arg 920 Glu Ile Gln Ser Ile Thr Asp Glu Ser Arg Gly Ser Ile Arg Arg Lys 930 935 Asn Pro Ala Asn Thr Arg Leu Arg Leu Asn Val Pro Glu Glu Thr Ala 950 955 945

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Gln Ser Ala Pro Ser Cys Pro Ser Ser Ser Pro Ser Pro Gly Glu Glu 980 985 990

Pro Glu Gly Glu Gly Glu Thr Asp Pro Glu Lys Val His Leu Thr Trp 995 1000 1005

Thr Lys Asp Lys Ser Val Ala Glu Lys Asn Lys Gly Pro Ser Pro Val 1010 1015 1020

Ser Ser Glu Gly Ile Lys Asp Phe Phe Ser Met Lys Pro Glu Trp Glu 1025 1030 1035 1040

Asn Leu Asn Gln Ser Asn Val Arg Arg Met His Thr Ala Val Arg Leu 1045 1050 1055

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Asn Met Pro Gly Pro Pro Arg Asn Arg Asn Gly Asp Glu Asn Tyr Met 1075 1080 1085

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ccc agg gaa a Pro Arg Glu A					
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aac ctg agc o Asn Leu Ser O 95					
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acc tgg att g Thr Trp Ile \			al Met Glu		
gcc atg tgc t Ala Met Cys 0 160		_	_		
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Leu Arg Gln His Lys Val Trp Arg Lys Cys Arg Met Arg Ile Phe Thr 865 870 875 880

Val Ala Gln Val Asp Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Gln 885 890 895

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<210> 129 <211> 76 <212> DNA <213> homo	sapiens					
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gtgtgtgtgt						76
<210> 130 <211> 21 <212> DNA <213> mouse	e					
<400× 130						

a)

21 tccagaaccg tggacagcgc C

<210> 131 <211> 1530 <212> DNA <213> mouse <220> <221> <222> <223> n=a,c, g, or t; sequence "nnnnn" comprises an undetermined number of nucleotides

<400> 131 qaattctqct actcattggc tttgtgtgcc taggcagtac ttttctctct tgggagttca 60 gtttttttgt ttttgttttt ttcatttacg aaatgaaagc tatacgttag ataagatgaa 120 180 cagttctgtc cagcaattct ccagggattt ttgaggactc aagctgggaa ccagggtgac tgtatagcta tttcctcttt ccattgctag tatgtttaga tctagatgca aagggtctaa 240 acctcaaatg tcactttctg aagcatctcc accattggcc caagtctggc ttcggggtcc 300 agggtaatca agctcctggg ggggtgctgg tttcagaaga ggaggacagg agagcatgtc 360 cggcctggaa gcctcagaaa gcaatcctgg attagaggat tagacattcc cgccccagga 420 gaaaatctgg tccccaaccg aggtggcctc attctcccct tccccactca cctggccagc 480 540 ttggggccag cataccetee ctagaggetg caccateteg ggtgggaagg aagtgetaca ggaccccagt gctccctgtt cctcgctgca tttcatccat agatcattcc acaggagaaa 600 tccccttgtc tgtggtgtgg actccagggt cttgtnnnnn gggccccgcc tgcgccagag 660 tgggggggc tgcgtctgct ctagggaagc gagggccgcg tcccccgcag catcccctcc 720 ataccacceg cgcaagcccc cagttttccc gtgaggggc caccggagct ctctgcccgc 780 ctctctgccc ttccctccct ccctccggtc ccccctcca aagaaaaccc gccagtggct 840 cacgcctcct gcataacggt atgaggtgag cagcgcccgc tactgagagg gggcgcgcgc 900 gggtgtgagc gtgtgtccgt gtgcgagtgt gtgtgcgccg ggcgggcggg cactgcagct 960 tcttcctccg tggagcggag agcaagcgag agagctcgag caagcgagcg agcggagaag 1020 gcgggcagag gggcgcgggc gaagcggcgc agccatcccg agcccggcgc cgcgcagcca 1080 ccangeteaa caacetgaeg gaetgegagg aeggegatgg gggageeaae eeeggtaage 1140 agtggtccgg gggcggcggg ggaggggta gcgaggagga aggaggagaa ggaggagaag 1200 gagaaggagg agggcagctt cggagtgggg ggcagagccg gggccgcctc ctgcagagga 1260 acagcctggg gctgaccggg gcggcggatg cctggaggct gcagggaggc tagctccatt 1320 ggaatgcgct aggcgctgtc cacggttctg gaatgcgccg cgctcccgag gcagattcgg 1380

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A.